09/686,497 STN SEARCH => file .nash => s galactosidase and common codon O FILE MEDLINE L1 2 FILE CAPLUS 1.3 0 FILE SCISEARCH O FILE LIFESCI L5 O FILE BIOSIS O FILE EMBASE L6 TOTAL FOR ALL FILES 2 GALACTOSIDASE AND COMMON CODON => d ibib abs ANSWER 1 OF 2 CAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2002:637845 CAPLUS DOCUMENT NUMBER: 137:180783 TITLE: Synthetic genes with optimized codon usage for recombinant protein expression in mammals INVENTOR(S): Seldon, Richard F.; Miller, Allan M.; Treco, Douglas S. PATENT ASSIGNEE(S): Transkaryotic Therapies, Inc., USA SOURCE: PCT Int. Appl., 115 pp. CODEN: PIXXD2 DOCUMENT TYPE: Patent LANGUAGE: English FAMILY ACC. NUM. COUNT: 1 PATENT INFORMATION: PATENT NO.

KIND DATE APPLICATION NO. DATE ----- ---- ---------A2 20020822 WO 2002064799 WO 2001-US42655 20011011 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG PRIORITY APPLN. INFO.: US 1999-407605 A1 19990929

US 2000-686497

A1 20001011

The present invention is directed to a synthetic nucleic acid sequence which encodes a protein wherein at least one non-common codon or less-common codon is replaced by a common codon. The synthetic nucleic acid sequence can include a continuous stretch of at least 90 codons all of which are common codons. Synthetic genes that have codon usages typical of highly expressed mammalian genes are prepd. for use in manuf. of the proteins using mammalian cell hosts. Recombinant expression of human Factor VIII and B-domain-deleted-FVIII, human Factor IX, and human .alpha.-galactosidase, in human fibroblast cells, is described.

=> d 2 ibib abs

ANSWER 2 OF 2 CAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 1989:434351 CAPLUS DOCUMENT NUMBER:

111:34351

TITLE: Codon usage determines translation rate in Escherichia

AUTHOR(S): Soerensen, Michael A.; Kurland, C. G.; Pedersen, Steen

CORPORATE SOURCE: Inst. Microbiol., Univ. Copenhagen, Copenhagen,

DK-1353, Den.

SOURCE: Journal of Molecular Biology (1989), 207(2), 365-77

CODEN: JMOBAK; ISSN: 0022-2836

DOCUMENT TYPE: Journal LANGUAGE: English

AB To det. whether differences in translation rate are correlated with

differences in codon usage or with differences in mRNA secondary structure, a small DNA fragment was inserted in the lacZ gene either directly or flanked by a few frame-shifting bases, leaving the reading frame of the lacZ gene unchanged. The fragment was chosen to have infrequent codons in 1 reading frame and common codons in the other. The insert in these constructs does not seem to give mRNAs that are able to form extensive secondary structures. The translation time for these modified lacZ mRNAs was measured with a reproducibility better than plus or minus 1 s. The mRNA with infrequent codons inserted has an .apprxeq.3-s longer translation time than the 1 with common codons. In another set of expts. 2 almost identical lacZ genes were constructed in which the lacZ mRNAs have the potential to generate stem structures with stabilities of .apprxeq.-75 kcal/mol. In this way it was possible to investigate the influence of mRNA structure on translation rate. This type of modified gene was generated in 2 reading frames with either common or infrequent codons similar to our 1st expts. The yield of protein from these mRNAs is reduced, probably due to the action in vivo of an RNase. Nevertheless, the data do not indicate that there is any effect of mRNA secondary structure on translation rate. In contrast, the data show that there is a difference in translation rate between infrequent codons and common codons that is of the order of 6-fold.

=> s common codon TOTAL FOR ALL FILES L14 78 COMMON CODON

=> s 114 and human TOTAL FOR ALL FILES

L21 16 L14 AND HUMAN

=> dup rem 121 PROCESSING COMPLETED FOR L21

L22 8 DUP REM L21 (8 DUPLICATES REMOVED)

=> d ibib abs

L22 ANSWER 1 OF 8 CAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2002:637845 CAPLUS

DOCUMENT NUMBER: 137:180783

TITLE: Synthetic genes with optimized codon usage for recombinant protein expression in mammals

INVENTOR(S): Seldon, Richard F.; Miller, Allan M.; Treco, Douglas

s.

PATENT ASSIGNEE(S): Transkaryotic Therapies, Inc., USA

SOURCE: PCT Int. Appl., 115 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

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PATENT NO. KIND DATE
                                               APPLICATION NO. DATE
     PATENT NO.
                                                -----
     WO 2002064799 A2 20020822
                                               WO 2001-US42655 20011011
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
              CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
              GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
              LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL,
              PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
              DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
PRIORITY APPLN. INFO.:
                                             US 1999-407605 A1 19990929
                                             US 2000-686497
                                                               Al 20001011
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AB The present invention is directed to a synthetic nucleic acid sequence which encodes a protein wherein at least one non-common codon or less-common codon is replaced by a common codon. The synthetic nucleic acid sequence can

include a continuous stretch of at least 90 codons all of which are common codons. Synthetic genes that have codon usages typical of highly expressed mammalian genes are prepd. for use in manuf. of the proteins using mammalian cell hosts. Recombinant expression of human Factor VIII and B-domain-deleted-FVIII, human Factor IX, and human .alpha.-galactosidase, in human fibroblast cells, is described. => s galactosidase and common codon O FILE MEDLINE L1 2 FILE CAPLUS L2 0 FILE SCISEARCH 1.3 O FILE LIFESCI 1.4 O FILE BIOSIS O FILE EMBASE 1.6 TOTAL FOR ALL FILES 2 GALACTOSIDASE AND COMMON CODON => d ibib abs ANSWER 1 OF 2 CAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2002:637845 CAPLUS DOCUMENT NUMBER: 137:180783 TITLE: Synthetic genes with optimized codon usage for recombinant protein expression in mammals Seldon, Richard F.; Miller, Allan M.; Treco, Douglas INVENTOR (S): Transkaryotic Therapies, Inc., USA PATENT ASSIGNEE(S): SOURCE: PCT Int. Appl., 115 pp. CODEN: PIXXD2 DOCUMENT TYPE: Patent LANGUAGE: English FAMILY ACC. NUM. COUNT: 1 PATENT INFORMATION: PATENT NO. KIND DATE APPLICATION NO. DATE _____

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PATENT NO. KIND DATE APPLICATION NO. DATE

WO 2002064799 A2 20020822 WO 2001-US42655 20011011

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM

RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

PRIORITY APPLN. INFO::

US 1999-407605 A1 19990929

US 2000-686497 A1 20001011
```

AB The present invention is directed to a synthetic nucleic acid sequence which encodes a protein wherein at least one non-common codon or less-common codon is replaced by a common codon. The synthetic nucleic acid sequence can include a continuous stretch of at least 90 codons all of which are common codons. Synthetic genes that have codon usages typical of highly expressed mammalian genes are prepd. for use in manuf. of the proteins using mammalian cell hosts. Recombinant expression of human Factor VIII and B-domain-deleted-FVIII, human Factor IX, and human .alpha.-galactosidase, in human fibroblast cells, is described.

=> d 2 ibib abs

L7 ANSWER 2 OF 2 CAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 1989:434351 CAPLUS

DOCUMENT NUMBER: 111:34351

TITLE: Codon usage determines translation rate in Escherichia

coli

AUTHOR(S): Soerensen, Michael A.; Kurland, C. G.; Pedersen, Steen

CORPORATE SOURCE: Inst. Microbiol., Univ. Copenhagen, Copenhagen,

DK-1353, Den.

Journal of Molecular Biology (1989), 207(2), 365-77 SOURCE:

CODEN: JMOBAK; ISSN: 0022-2836

DOCUMENT TYPE: LANGUAGE:

Journal Enalish

To det. whether differences in translation rate are correlated with differences in codon usage or with differences in mRNA secondary structure, a small DNA fragment was inserted in the lacZ gene either directly or flanked by a few frame-shifting bases, leaving the reading frame of the lacZ gene unchanged. The fragment was chosen to have infrequent codons in 1 reading frame and common codons in the other. The insert in these constructs does not seem to give mRNAs that are able to form extensive secondary structures. The translation time for these modified lacZ mRNAs was measured with a reproducibility better than plus or minus 1 s. The mRNA with infrequent codons inserted has an .apprxeq.3-s longer translation time than the 1 with ${\color{red}\mathbf{common}}$ codons. In another set of expts. 2 almost identical lacZ genes were constructed in which the lacZ mRNAs have the potential to generate stem structures with stabilities of .apprxeq.-75 kcal/mol. In this way it was possible to investigate the influence of mRNA structure on translation rate. This type of modified gene was generated in 2 reading frames with either common or infrequent codons similar to our 1st expts. The yield of protein from these mRNAs is reduced, probably due to the action in vivo of an RNase. Nevertheless, the data do not indicate that there is any effect of mRNA secondary structure on translation rate. In contrast, the data show that there is a difference in translation rate between infrequent codons and common codons that is of the order of

=> s common codon TOTAL FOR ALL FILES 78 COMMON CODON L14

6-fold.

=> s 114 and human TOTAL FOR ALL FILES

16 L14 AND HUMAN

=> dup rem 121

PROCESSING COMPLETED FOR L21

8 DUP REM L21 (8 DUPLICATES REMOVED)

=> d ibib abs

L22 ANSWER 1 OF 8 CAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2002:637845 CAPLUS

DOCUMENT NUMBER: 137:180783

TITLE: Synthetic genes with optimized codon usage for

recombinant protein expression in mammals

INVENTOR(S): Seldon, Richard F.; Miller, Allan M.; Treco, Douglas

s.

PATENT ASSIGNEE(S): Transkaryotic Therapies, Inc., USA PCT Int. Appl., 115 pp.

SOURCE:

CODEN: PIXXD2 DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PAT	TENT 1	NO.		KI	ND	DATE			A	PPLI	CATI	N NC	o. :	DATE			
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WO	2002	0647	99	A:	2	2002	0822		W	20	01-U	S426	55 :	2001	1011		
	W:	ΑE,	AG,	AL,	AM,	ΑT,	ΑU,	AZ,	BA,	BB,	BG,	BR,	BY,	ΒZ,	CA,	CH,	CN,
		CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FI,	GB,	GD,	GE,	GH,
		GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	ΚP,	KR,	ΚZ,	LC,	LK,	LR,
		LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NO,	NZ,	PH,	PL,
		PT,	RO,	RU,	SD,	SE,	SG,	SI,	SK,	SL,	TJ,	TM,	TR,	TT,	TZ,	UA,	UG,
		US,	UZ,	VN,	YU,	ZA,	ZW,	AM,	AZ,	BY,	KG,	ΚZ,	MD,	RU,	ТJ,	TM	
	RW:	GH,	GM,	ΚE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	UG,	ZW,	AT,	BE,	CH,	CY,
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		ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN,	GQ,	GW,	ML,	MR,	NE,	SN,	TD,	TG	
ORITY	APP	LN.	INFO.	. :				i	JS 1:	999-	4076	05	A1 :	19990	0929		

US 2000-686497 Al 20001011

The present invention is directed to a synthetic nucleic acid sequence which encodes a protein wherein at least one non-common codon or less-common codon is replaced by a common codon. The synthetic nucleic acid sequence can include a continuous stretch of at least 90 codons all of which are common codons. Synthetic genes that have codon usages typical of highly expressed mammalian genes are prepd. for use in manuf. of the proteins using mammalian cell hosts. Recombinant expression of human Factor VIII and B-domain-deleted-FVIII, human Factor IX, and human .alpha.-galactosidase, in human fibroblast cells, is described.

=> s common codon TOTAL FOR ALL FILES 78 COMMON CODON

=> s 17 and human TOTAL FOR ALL FILES 16 L7 AND HUMAN

=> dup rem 114 PROCESSING COMPLETED FOR L14 8 DUP REM L14 (8 DUPLICATES REMOVED)

=> d ibib abs

L15 ANSWER 1 OF 8 CAPLUS COPYRIGHT 2002 ACS 2002:637845 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 137:180783

TITLE: Synthetic genes with optimized codon usage for

recombinant protein expression in mammals

INVENTOR(S): Seldon, Richard F.; Miller, Allan M.; Treco, Douglas

PATENT ASSIGNEE(S): Transkaryotic Therapies, Inc., USA

PCT Int. Appl., 115 pp. SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

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KIND DATE
                                                 APPLICATION NO. DATE
     PATENT NO.
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                         A2 20020822
                                                WO 2001-US42655 20011011
     WO 2002064799
          W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
              CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
               LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL,
               PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
          RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
               DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
               BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                                              US 1999-407605 A1 19990929
US 2000-686497 A1 20001011
PRIORITY APPLN. INFO.:
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The present invention is directed to a synthetic nucleic acid sequence AB which encodes a protein wherein at least one non-common codon or less-common codon is replaced by a common codon. The synthetic nucleic acid sequence can include a continuous stretch of at least 90 codons all of which are common codons. Synthetic genes that have codon usages typical of highly expressed mammalian genes are prepd. for use in manuf. of the proteins using mammalian cell hosts. Recombinant expression of human Factor VIII and B-domain-deleted-FVIII, human Factor IX, and human .alpha.-galactosidase, in human fibroblast cells, is described.

L15 ANSWER 2 OF 8 MEDLINE DUPLICATE 1

ACCESSION NUMBER: 1998025109 MEDLINE

DOCUMENT NUMBER: 98025109 PubMed ID: 9360634

TITLE: A nucleotide polymorphism in ERCC1 in human ovarian cancer cell lines and tumor tissues.

AUTHOR: Yu J J; Mu C; Lee K B; Okamoto A; Reed E L; Bostick-Bruton

F; Mitchell K C; Reed E

CORPORATE SOURCE: Developmental Therapeutics Department, National Cancer

Institute, Bethesda, MD 20892, USA.

SOURCE: MUTATION RESEARCH, (1997 Sep) 382 (1-2) 13-20.

Journal code: 0400763. ISSN: 0027-5107.

PUB. COUNTRY: Netherlands

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals OTHER SOURCE: GENBANK-AF001925

ENTRY MONTH: 199712

ENTRY DATE: Entered STN: 19980109

Last Updated on STN: 19980109 Entered Medline: 19971201

We studied the DNA sequence of the entire coding region of ERCC1 gene, in AR five cell lines established from human ovarian cancer (A2780, A2780/CP70, MCAS, OVCAR-3, SK-OV-3), 29 human ovarian cancer tumor tissue specimens, one human T-lymphocyte cell line (H9), and non-malignant human ovary tissue (NHO). Samples were assayed by PCR-SSCP and DNA sequence analyses. A silent mutation at codon 118 (site for restriction endonuclease MaeII) in exon 4 of the gene was detected in MCAS, OVCAR-3 and SK-OV-3 cells, and NHO. This mutation was a C-->T transition, that codes for the same amino acid: asparagine. This transition converts a common codon usage (AAC) to an infrequent codon usage (AAT), whereas frequency of use is reduced two-fold. This base change was associated with a detectable band shift on SSCP analysis. For the 29 ovarian cancer specimens, the same base change was observed in 15 tumor samples and was associated with the same band shift in exon 4. Cells and tumor tissue specimens that did not contain the C-->T transition, did not show the band shift in exon 4. Our data suggest that this alteration at codon 118 within the ERCC1 gene, may exist in platinum-sensitive and platinum-resistant ovarian cancer tissues.

L15 ANSWER 3 OF 8 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.

ACCESSION NUMBER: 1998275804 EMBASE

TITLE: A nucleotide polymorphism in ERCC1 in human

ovarian cancer cell lines and tumor tissues.

AUTHOR: Jing Jie Yu; Mu C.; Kang Bo Lee; Okamoto A.; Reed E.L.;

Bostick-Bruton F.; Mitchell K.C.; Reed E.

CORPORATE SOURCE: E. Reed, Medical Ovarian Cancer Section, Devtl.

Therapeutics Department, National Cancer Institute,

Bethesda, MD 20892, United States. reed92@helix.nih.gov Mutation Research - Mutation Research Genomics, (1997)

SOURCE: Mutation Research 382/1-2 (13-20).

Refs: 29

ISSN: 1383-5726 CODEN: MMRGFK

PUBLISHER IDENT.: S 1383-5726(97)00004-6

COUNTRY: Netherlands
DOCUMENT TYPE: Journal; Article

FILE SEGMENT: 010 Obstetrics and Gynecology

016 Cancer 022 Human Genetics

037 Drug Literature Index

LANGUAGE: English SUMMARY LANGUAGE: English

AB We studied the DNA sequence of the entire coding region of ERCC1 gene, in five cell lines established from human ovarian cancer (A2780, A2780/CP70, MCAS, OVCAR-3, SK-OV-3), 29 human ovarian cancer tumor tissue specimens, one human T-lymphocyte cell line (H9), and non-malignant human ovary tissue (NHO). Samples were assayed by PCR-SSCP and DNA sequence analyses. A silent mutation at codon 118 (site for restriction endonuclease MaeII) in exon 4 of the gene was detected in MCAS, OVCAR-3 and SK-OV-3 cells, and NHO. This mutation was a C .fwdarw. T transition, that codes for the same amino acid: asparagine. This transition converts a common codon usage (AAC) to

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L15 ANSWER 4 OF 8 MEDLINE

ACCESSION NUMBER: 97179330 MEDLINE

DOCUMENT NUMBER: 97179330 PubMed ID: 9027616

TITLE: Clinical detection of lung cancer progression markers.

AUTHOR: Tockman M S

CORPORATE SOURCE: Johns Hopkins University School of Hygiene and Public

Health, Department of Environmental Health Sciences,

Baltimore, Maryland 21205, USA.

CONTRACT NUMBER: 1P50 CA58184-01 (NCI)

SOURCE: JOURNAL OF CELLULAR BIOCHEMISTRY. SUPPLEMENT, (1996) 25

177-84. Ref: 36

Journal code: 8207539. ISSN: 0733-1959.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

General Review; (REVIEW)

(REVIEW, TUTORIAL)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199704

ENTRY DATE: Entered STN: 19970424

Last Updated on STN: 19970424 Entered Medline: 19970417

Lung cancer is the leading cause of cancer-related deaths in western countries. The prognosis for patients with lung cancer depends primarily on the stage of the tumor at the time of clinical diagnosis. New understanding of tumor biology has turned attention away from detection of clinical lung cancer, usually metastatic at presentation, toward recognition of genetic and protein markers which precede malignancy. Mutations of four types of genes contribute to the process of epithelial carcinogenesis by modifying control of cell growth. Examples of three of these changes have been detected in pre-malignant sputum, and validated in subsequent tumor. We have identified gene products (tumor associated and differentiation protein antigens), mutations of k-ras and p53, and microsatellite alterations as potential markers of subsequent malignancy. We consider the morphologic progression seen in archived sputum cells as the paradigm of neoplastic development in the lung. Although the NCl collaborative trials had shown that this progression is not recognized sufficiently often (sensitive) to be useful for lung cancer screening, this progression may be used to assess the timing of gene and peptide markers of carcinogenesis. Previous work has shown that at the time Johns Hopkins Lung Project sputum cells express moderately atypical metaplasia, 53% (8/15) of sputum specimens expressed ${\bf common}$ (${\bf codon}$ 12) k-ras or (codons 273 or 281) p53 mutations. Other investigators have reported that earlier morphologic changes (metaplasia) accompany 3p and 9p losses of heterozygosity. These observations suggest that 3p and 9p loss likely precede k-ras or p53 mutations. Our preliminary data demonstrate that over-expression of a 31 kD tumor associated antigen recently purified, sequenced, and identified as heterogeneous nuclear ribonucleoprotein (hnRNP) A2 (with cross reactivity to splice variant B1), is expressed in most lung cancer cases before any morphologic abnormality. Comparison of the accuracy of this marker with sputum cytology will determine its value for early lung cancer detection. Preliminary evidence confirms this marker greatly improves the accuracy of standard sputum cytology for detection of lung carcinogenesis. Clinical intervention trials must be undertaken to determine whether modulation of hnRNP overexpression is useful as an intermediate endpoint for chemoprevention.

L15 ANSWER 5 OF 8 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.

ACCESSION NUMBER: 97054548 EMBASE

DOCUMENT NUMBER: 1997054548

TITLE: Clinical detection of lung cancer progression markers.

AUTHOR: Tockman M.S.

CORPORATE SOURCE: M.S. Tockman, School of Hygiene and Public Health, Johns

Hopkins University, 615 N. Wolfe Street, Baltimore, MD

21205, United States

SOURCE: Journal of Cellular Biochemistry, (1996) 63/SUPPL. 25

(177-184). Refs: 35

ISSN: 0730-2312 CODEN: JCEBD5

COUNTRY: United States
DOCUMENT TYPE: Journal; Article
FILE SEGMENT: 016 Cancer

029 Clinical Biochemistry

LANGUAGE: English SUMMARY LANGUAGE: English

Lung cancer is the leading cause of cancer-related deaths in western countries. The prognosis for patients with lung cancer depends primarily on the stage of the tumor at the time of clinical diagnosis. New understanding of tumor biology has turned attention away from detection of clinical lung cancer, usually metastatic at presentation, toward recognition of genetic and protein markers which precede malignancy. Mutations of four types of genes contribute to the process of epithelial carcinogenesis by modifying control of cell growth. Examples of three of these changes have been detected in pre- malignant sputum, and validated in subsequent tumor. We have identified gene products (tumor associated and differentiation protein antigens), mutations of k-ras and p53, and microsatellite alterations as potential markers of subsequent malignancy. We consider the morphologic progression seen in archived sputum cells as the paradigm of neoplastic development in the lung. Although the NCI collaborative trials had shown that this progression is not recognized sufficiently often (sensitive) to be useful for lung cancer screening, this progression may be used to assess the timing of gene and peptide markers of carcinogenesis. Previous work has shown that at the time Johns Hopkins Lung Project sputum cells express moderately atypical metaplasia, 53% (8/15) of sputum specimens expressed common (codon 12) k-ras or (codons 273 or 281) p53 mutations. Other investigators have reported that earlier morphologic changes (metaplasia) accompany 3p and 9p losses of heterozygosity. These observations suggest that 3p and 9p loss likely precede k-ras or p53 mutations. Our preliminary data demonstrate that over-expression of a 31 kD tumor associated antigen recently purified, sequenced, and identified as heterogeneous nuclear ribonucleoprotein (hnRNP) A2 (with cross reactivity to splice variant B1), is expressed in most lung cancer cases before any morphologic abnormality. Comparison of the accuracy of this marker with sputum cytology will determine its value for early lung cancer detection. Preliminary evidence confirms this marker greatly improves the accuracy of standard sputum cytology for detection of lung carcinogenesis. Clinical intervention trials must be undertaken to determine whether modulation of hnRNP

L15 ANSWER 6 OF 8 MEDLINE DUPLICATE 2

ACCESSION NUMBER: 97083381 MEDLINE

DOCUMENT NUMBER: 97083381 PubMed ID: 8929955

TITLE: Familial adenomatous polyposis in a 5 year old child: a clinical, pathological, and molecular genetic study.

overexpression is useful as an intermediate endpoint for chemoprevention.

AUTHOR: Distante S; Nasioulas S; Somers G R; Cameron D J; Young M A; Forrest S M; Gardner R J

CORPORATE SOURCE: The Murdoch Institute, Royal Children's Hospital,

Melbourne, Australia.

SOURCE: JOURNAL OF MEDICAL GENETICS, (1996 Feb) 33 (2) 157-60.

Journal code: 2985087R. ISSN: 0022-2593.

PUB. COUNTRY: ENGLAND: United Kingdom

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH:

199705

ENTRY DATE: Entered STN: 19970523

Last Updated on STN: 19970523

Entered Medline: 19970512

AB A girl aged 5 years 8 months presented with rectal bleeding; her father had had familial adenomatous polyposis (FAP) and a colectomy at the age of 23. Endoscopy showed extensive polyposis and she had a colectomy. The proband and her father had the **common codon** 1309 5 bp

deletion APC mutation. This mutation predisposes to early onset of FAP, and consideration needs to be given to having molecular testing of at risk members of these families done in childhood.

L15 ANSWER 7 OF 8 CAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 1995:66454 CAPLUS

DOCUMENT NUMBER: 122:78365

TITLE: An A-to-C substitution involving the translation initiation codon in a patient with myophosphorylase

deficiency (McArdle's disease)

AUTHOR(S): Tsujino, Seiichi; Rubin, Laurence A.; Shanske, Sara;

DiMauro, Salvatore

CORPORATE SOURCE: H. Houston Merritt Clinical Research Center Muscular

Dystrophy and related diseases, New York, NY, 10032,

USA

SOURCE: Human Mutation (1994), 4(1), 73-5 CODEN: HUMUE3; ISSN: 1059-7794

DOCUMENT TYPE: Journal LANGUAGE: English

AB RFLP screening of white blood cell genomic DNA from a 36-yr-old woman with McArdle's disease indicated that she had the relatively common codon 49 nonsense mutation in one allele but another mutation in her other allele. Sequencing of the other allele revealed an A-to-C substitution in the initiation codon for the myophosphorylase gene, changing it to CTG.

L15 ANSWER 8 OF 8 MEDLINE DUPLICATE 3

ACCESSION NUMBER: 89255401 MEDLINE

DOCUMENT NUMBER: 89255401 PubMed ID: 2656695

TITLE: Expression of human thymidylate synthase in

Escherichia coli.

COMMENT: Erratum in: J Biol Chem 1994 Dec 2;269(48):30740

AUTHOR: Davisson V J; Sirawaraporn W; Santi D V

CORPORATE SOURCE: Department of Biochemistry and Biophysics, University of

California, San Francisco 94143.

CONTRACT NUMBER: 5T32CA09270 (NCI)

CA14394 (NCI)

SOURCE: JOURNAL OF BIOLOGICAL CHEMISTRY, (1989 Jun 5) 264 (16)

9145-8.

Journal code: 2985121R. ISSN: 0021-9258.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 198907

ENTRY DATE: Entered STN: 19900306

Last Updated on STN: 19980206 Entered Medline: 19890705

AB A cDNA clone encoding thymidylate synthase (TS) has been isolated from a human T-cell library and modified in the 5'-untranslated region to incorporate several unique cloning sites. The gene has been cloned as a cassette into several Escherichia coli expression vectors which did not provide detectable amounts of the enzyme. A successful approach used a constitutive E. coli expression vector developed for the enzyme from Lactobacillus casei. A 115-base pair 5'-untranslated region from the L. casei TS which contains a ribosomal binding site and other regulatory sequences has been fused to the coding region of the $\boldsymbol{human}\ TS$ gene to provide a construct that is expressed in E. coli. The level of expression was further enhanced by altering the nucleotide sequence of the first 90 base pairs to accommodate common codon use in E. coli. In our best expression system, catalytically active human TS is expressed to a level that represents about 1.6% of the total soluble protein. The recombinant human TS has been purified and characterized; except for the presence of an amino-terminal blocking group, the enzyme has physical and kinetic properties similar to the enzyme isolated from human cells.

=> s galactosidase and fabry TOTAL FOR ALL FILES 2081 GALACTOSIDASE AND FABRY => s 12 and human TOTAL FOR ALL FILES 1585 L2 AND HUMAN => s 17 and human TOTAL FOR ALL FILES 1585 L7 AND HUMAN \Rightarrow s 121 and .alpha. TOTAL FOR ALL FILES 1491 L21 AND .ALPHA. => s 128 not 1990-2002/py 175 FILE MEDLINE L29 57 FILE CAPLUS L30 4 FILE SCISEARCH L31 3 FILE LIFESCI L32 107 FILE BIOSIS 1.33 T.34 98 FILE EMBASE TOTAL FOR ALL FILES 444 L28 NOT 1990-2002/PY => d 1-10 129 ibib abs YOU HAVE REQUESTED DATA FROM FILE 'MEDLINE' - CONTINUE? (Y)/N:y L29 ANSWER 1 OF 175 MEDLINE ACCESSION NUMBER: 91027226 MEDLINE DOCUMENT NUMBER: 91027226 PubMed ID: 2908672 TITLE: Anderson-Fabry disease--family linkage studies using two polymorphic X-linked DNA probes. AUTHOR: Morgan S H; Cheshire J K; Wilson T M; MacDermot K; Crawfurd M.A CORPORATE SOURCE: Division of Inherited Metabolic Diseases, MRC Clinical Research Centre, Harrow, Middlesex, UK. PEDIATRIC NEPHROLOGY, (1987 Jul) 1 (3) 536-9. SOURCE: Journal code: 8708728. ISSN: 0931-041X. PUB. COUNTRY: GERMANY: Germany, Federal Republic of DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE) LANGUAGE: English FILE SEGMENT: Priority Journals ENTRY MONTH: 199012 ENTRY DATE: Entered STN: 19910208 Last Updated on STN: 19950206 Entered Medline: 19901227 Anderson-Fabry disease is an X-linked lysosomal storage disorder due to alpha-galactosidase A deficiency. In affected males there is a high mortality in early adult life due to renal failure and cardiovascular complications. We describe our preliminary results from genetic linkage studies in five families using two polymorphic DNA probes, DXS17 and DXYS1, mapping to an area on the long arm of the ${\tt X}$ chromosome between Xq13-22. DXS17 identified a Taql polymorphism closely linked to the disease locus in three families (lodmax Z = 4.23. at a recombination fraction decreases theta = 0.0). Restriction fragment length polymorphisms detected by DXYS1 were not linked. L29 ANSWER 2 OF 175 MEDLINE

ACCESSION NUMBER: 90299318 MEDLINE

DOCUMENT NUMBER: 90299318 PubMed ID: 2561653

TITLE: [Fabry's disease: kidney insufficiency in

heterozygous patient].

Malattia di Fabry: insufficienza renale in

eterozigote.

AUTHOR: Pravata G; Pinto G; Noto G; Arico M

SOURCE: GIORNALE ITALIANO DI DERMATOLOGIA E VENEREOLOGIA, (1989

Nov-Dec) 124 (11-12) 505-9.

Journal code: 8102852. ISSN: 0026-4741.

PUB. COUNTRY: Italy

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: Italian

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199008

ENTRY DATE: Entered STN: 19900907

Last Updated on STN: 19990129 Entered Medline: 19900806

We report a rare heterozygous status for Fabry's gene with severe kidney involvement and normal alpha-galactosidase

A activity, together with the intrafamilial variations in the clinical expression of the disease. The random X inactivation hypothesis seems to

explain such a variable expression of the alpha-

galactosidase gene in our cases.

L29 ANSWER 3 OF 175 MEDLINE

ACCESSION NUMBER: 90296528 MEDLINE

DOCUMENT NUMBER: 90296528 PubMed ID: 2561643

TITLE: The gene encoding alpha-galactosidase A

and gene rearrangements causing Fabry disease.

AUTHOR: Kornreich R; Bishop D.F; Desnick R J

Division of Medical and Molecular Genetics, Mount Sinai CORPORATE SOURCE:

School of Medicine, New York, NY 10029.

CONTRACT NUMBER: 2 T32 HD07105 (NICHD)

5 M01 RR00071 (NCRR)

5 R01 DK34045 (NIDDK) ---

TRANSACTIONS OF THE ASSOCIATION OF AMERICAN PHYSICIANS, SOURCE:

(1989) 102 30-43.

Journal code: 7506109. ISSN: 0066-9458.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals 199007

ENTRY MONTH:

ENTRY DATE: Entered STN: 19900907

> Last Updated on STN: 19990129 Entered Medline: 19900727

L29 ANSWER 4 OF 175 MEDLINE

ACCESSION NUMBER: 90145947 MEDLINE

DOCUMENT NUMBER: 90145947 PubMed ID: 2559640

TITLE: [Fabry's disease and Klippel-Trenaunay syndrome

of the 4 limbs].

Maladie de Fabry et syndrome de Klippel-Trenaunay

des quatre membres. AUTHOR: Enjolras O; Leibowitch M; Riche M C; Lizop M; Escande J P

CORPORATE SOURCE: Service de Dermatologie, Hopital Tarnier, Paris.

SOURCE:

ANNALES DE DERMATOLOGIE ET DE VENEREOLOGIE, (1989) 116 (11)

788-90.

Journal code: 7702013. ISSN: 0151-9638.

PUB. COUNTRY: France

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: French

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199003

Entered STN: 19900328 ENTRY DATE:

Last Updated on STN: 19900328 Entered Medline: 19900315

L29 ANSWER 5 OF 175 MEDLINE

ACCESSION NUMBER: 90116512 MEDLINE

DOCUMENT NUMBER: 90116512 PubMed ID: 2855953

TITLE: Alpha-galactosidase A deficiency--

Fabry's disease.

AUTHOR: Tsuji S

SOURCE: TANPAKUSHITSU KAKUSAN KOSO. PROTEIN, NUCLEIC ACID, ENZYME, (1988 Apr) 33 (5) 745-8.

Journal code: 0413762. ISSN: 0039-9450.

PUB. COUNTRY:

Japan

Journal; Article; (JOURNAL ARTICLE) DOCUMENT TYPE:

LANGUAGE:

Japanese

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

199002

ENTRY DATE:

Entered STN: 19900328

Last Updated on STN: 19990129 Entered Medline: 19900220

L29 ANSWER 6 OF 175 MEDLINE

ACCESSION NUMBER:

90080835 MEDLINE

DOCUMENT NUMBER:

90080835 PubMed ID: 2556612

TITLE:

Detection of Fabry's disease carriers by enzyme

assay of hair roots.

COMMENT: AUTHOR:

Comment in: J Inherit Metab Dis. 1989;12(4):491-2 Hatton C E; Cooper A; Sardharwalla I B

CORPORATE SOURCE:

Willink Biochemical Genetics Unit, Royal Manchester

Children's Hospital, Pendleburg, UK.

SOURCE:

JOURNAL OF INHERITED METABOLIC DISEASE, (1989) 12 Suppl 2

369-71

PUB. COUNTRY:

Journal code: 7910918. ISSN: 0141-8955. Netherlands

DOCUMENT TYPE:

LANGUAGE:

Journal; Article; (JOURNAL ARTICLE)

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

199001

ENTRY DATE:

Entered STN: 19900328

Last Updated on STN: 19990129 Entered Medline: 19900125

L29 ANSWER 7 OF 175 MEDLINE

ACCESSION NUMBER: 90068337

MEDLINE

DOCUMENT NUMBER:

90068337 PubMed ID: 2555802 Anderson Fabry disease -- an identifiable disorder.

TITLE: AUTHOR: SOURCE:

Wakeel R; Shibib K; Chapman R; Dunnigan M PRACTITIONER, (1989 Mar 8) 233 (1464) 294-5.

Journal code: 0404245. ISSN: 0032-6518.

PUB. COUNTRY:

ENGLAND: United Kingdom

DOCUMENT TYPE: LANGUAGE:

Journal; Article; (JOURNAL ARTICLE)

English

Priority Journals FILE SEGMENT:

ENTRY MONTH:

199001

Entered STN: 19900328 ENTRY DATE:

Last Updated on STN: 19990129 Entered Medline: 19900102

L29 ANSWER 8 OF 175 MEDLINE

ACCESSION NUMBER: 89360761

MEDLINE

DOCUMENT NUMBER:

PubMed ID: 2504843 89360761

TITLE:

Transvenous permanent pacemaker implantation for

Fabry's disease. 3 cases report.

AUTHOR:

Yoshida K; Murase M; Maseki T; Usui A; Ina H; Abe T NIPPON KYOBU GEKA GAKKAI ZASSHI. JOURNAL OF THE JAPANESE

SOURCE:

ASSOCIATION FOR THORACIC SURGERY, (1989 Feb) 37 (2) 386-90. Journal code: 19130180R. ISSN: 0369-4739.

PUB. COUNTRY:

DOCUMENT TYPE:

Japan Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

Japanese

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

198909

ENTRY DATE:

Entered STN: 19900309

Last Updated on STN: 19990129 Entered Medline: 19890928

Three men with Fabry's disease (angiokeratoma corporis diffusum universal) are described. In the first patient, atrial fibrillation appeared, and a permanent cardiac pacemaker (VVI) was implanted. Sick sinus syndrome with complete atrioventricular block was occurred on the second patient. Transvenous pacemaker (DDD) implantation was performed for him. The last patient was younger brother of the second patient. He

demonstrated complete atrio-ventricular block, so cardiac pace maker (VAT) was implanted. They showed a low value of granulocyte's alphagalactosidase activity. During 1 to 4 year follow up period, they showed no trouble about pacemaking. Fabry's disease is an disorder of glycosphingolipid metabolism. This disorder is characterized by the accumulation of trihexosyl ceramide in many sites. Cardiac involvement and abnormal electrocardiographic manifestations are common in this disorder. Permanent cardiac pacemaker is necessary for severe bradycardia caused by this disorder.

L29 ANSWER 9 OF 175 MEDLINE

ACCESSION NUMBER: 89355456 MEDLINE

PubMed ID: 2504516 DOCUMENT NUMBER: 89355456

TITLE: Angiokeratoma corporis diffusum in GM1 gangliosidosis, type

Beratis N G; Varvarigou-Frimas A; Beratis S; Sklower S L AUTHOR: Department of Pediatrics, University of Patras Medical

CORPORATE SOURCE:

School, Greece.

CLINICAL GENETICS, (1989 Jul) 36 (1) 59-64. SOURCE:

Journal code: 0253664. ISSN: 0009-9163.

PUB. COUNTRY: Denmark

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH:

198910

Entered STN: 19900309 ENTRY DATE:

Last Updated on STN: 19990129

Entered Medline: 19891004

A patient with severe deficiency of beta-galactosidase, who developed skin lesions of angiokeratoma corporis diffusum between the 3rd and 10th month of life, is described. The activity of other lysosomal enzymes, including alpha-neuraminidase, was normal. The first signs of the disease were noticed during the first month of life. By 3months coarseness of the face and psychomotor retardation were present. In addition to angiokeratoma, he had large mongolian spots and several scattered slate-blue spots of pigmentation over his body. With the exception of the skin lesions, the other clinical signs and the course of the psychomotor deterioration were within the clinical picture of GM1 gangliosidosis, Type 1. Angiokeratoma, a manifestation of several lysosomal disorders, may appear in GM1 gangliosidosis during the first year of life.

L29 ANSWER 10 OF 175 MEDLINE

ACCESSION NUMBER: 89323245 MEDLINE

DOCUMENT NUMBER: 89323245 PubMed ID: 2546612

TITLE: [Substrate specificity of multiple forms of human

alpha-D-galactosidase and alpha

-D-fucosidase].

Substratnaia spetsifichnost' mnozhestvennykh form

alpha-D-galaktozidazy i alpha

-D-fukozidazy cheloveka.

AUTHOR: Baskaeva E M; Shono N I; Kozlova I K; Vidershain G Ia

SOURCE: BIOKHIMIIA, (1989 Mar) 54 (3) 421-6. Journal code: 0372667. ISSN: 0320-9725.

PUB. COUNTRY: USSR

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: Russian

FILE SEGMENT: Priority Journals

ENTRY MONTH: 198908

ENTRY DATE: Entered STN: 19900309

Last Updated on STN: 20000303 Entered Medline: 19890829

It was shown that human alpha-D-galactosidase is represented by multiple forms, only one of which can also split

alpha-D-fucoside. Fabry's disease was found to be associated not only with the deficiency of the alpha-D-

galactosidase total activity but also with the deficiency of the alpha-D-fucosidase activity. The decrease in the alpha

-D-galactosidase activity is due to the lack of two enzyme forms, while the profile of alpha-D-fucosidase multiple forms during isoelectric focusing of human enzyme preparations is

modified very little in comparison with the normal one. The deficiency of both enzymes was expressed in most degree in leukocytes as compared to other tissues. The residual activities of alpha-D-galactosidase and alpha-D-fucosidase in leukocytes were equal to 3.5 and 21%, respectively. Since the decrease in the alpha-D-fucosidase activity was not so noticeable as in the alpha-D-galactosidase activity, it may be expected that the determination of the alpha-D-fucosidase activity can no longer be regarded as a reliable test for the diagnosis of Fabry 's disease. The data obtained suggest that alpha-D-galactoside and alpha-D-fucoside are split by the same enzyme, the multiple forms of which are characterized by selective specificity towards these substrates.

WEST Search History

DATE: Monday, October 28, 2002

Set Name side by side	Query	Hit Count	Set Name result set
•	SPT,PGPB,JPAB,EPAB,DWPI; THES=ASSIGNEE; PLUR=YES;		
L10	common codon	25	L10
L9	galactosidase and common adj1 codon	4	L9
L8	human and galactosidase and (nucleic acid or dna or cdna or rna or mrna) and common adj1 codon	4	L8
L7	human and galactosidase and (nucleic acid or dna or cdna or rna of mrna) and common adj1 codon	4	L7
DB=US	PT,PGPB; THES=ASSIGNEE; PLUR=YES; OP=ADJ		
L6	human same galactosidase and (nucleic acid or dna or cdna or rna of mrna) and common adj1 codon	0	L6
L5	L4 and common codon	0	L5
L4	human same galactosidase and (nucleic acid or dna or cdna or rna of mrna)	2437	L4
L3	L2 and (nucleic acid or dna or cdna or rna of mrna)	14694	L3
L2	galactosidase	16508	L2
DB=US	PT; THES=ASSIGNEE; PLUR=YES; OP=ADJ		
L1	galactosidase	13369	L1

END OF SEARCH HISTORY

WEST

Generate Collection

Print

Search Results - Record(s) 1 through 4 of 4 returned.

1. Document ID: US 20020123083 A1

L9: Entry 1 of 4

File: PGPB

Sep 5, 2002

PGPUB-DOCUMENT-NUMBER: 20020123083

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20020123083 A1

TITLE: Nucleic acid endocing growth factor protein

PUBLICATION-DATE: September 5, 2002

INVENTOR-INFORMATION:

NAME

CITY

STATE

COUNTRY RU

RULE-47

Shigeta, Ron T. JR.

Berkeley

CA

US

Siani-Rose, Michael A.

San Francisco

ncisco CA US

US-CL-CURRENT: 435/7.23; 435/320.1, 435/325, 435/69.4, 530/399, 536/23.5, 702/19, 800/8

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KMC
Draws D	eso li	nage									

2. Document ID: US 6232458 B1

L9: Entry 2 of 4

File: USPT

May 15, 2001

US-PAT-NO: 6232458

DOCUMENT-IDENTIFIER: US 6232458 B1

TITLE: Synthetic polynucleotides encoding tropoelastin

DATE-ISSUED: May 15, 2001

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Weiss; Anthony Steven Sydney AU
Martin; Stephen Lewis Sedgley GB

US-CL-CURRENT: $\underline{536}/\underline{23.5}$; $\underline{435}/\underline{252.33}$, $\underline{435}/\underline{254.1}$, $\underline{435}/\underline{254.2}$, $\underline{435}/\underline{320.1}$, $\underline{435}/\underline{69.1}$, $\underline{435}/\underline{69.7}$, $\underline{530}/\underline{353}$, $\underline{536}/\underline{23.4}$, $\underline{536}/\underline{24.1}$, $\underline{536}/\underline{24.2}$

ABSTRACT:

Recombinant tropoelastins and variants of recombinant tropoelastins produced from synthetic polynucleotides, as well as the synthetic polynucleotides themselves are provided. Also provided are cross-linked elastins or elastin-like products prepared from the tropoelastins or variants.

24 Claims, 21 Drawing figures

Exemplary Claim Number: 1 Number of Drawing Sheets: 21

Full | Title | Citation | Front | Review | Classification | Date | Reference | Sequences | Attachments | KW.

3. Document ID: US 5955277 A

L9: Entry 3 of 4

File: USPT

Sep 21, 1999

US-PAT-NO: 5955277

DOCUMENT-IDENTIFIER: US 5955277 A

TITLE: Mutant cDNA encoding the p85.alpha. subunit of phosphatidylinositol 3-kinase

DATE-ISSUED: September 21, 1999

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Hansen; Torben Hellerup DK

Andersen; Carsten Bo Los Altos CA

Pedersen; Oluf Borbye Holte DK

US-CL-CURRENT: 435/6; 435/91.2, 536/23.1

ABSTRACT:

The present invention relates to a mutant cDNA sequence encoding the regulatory p85.alpha. subunit of phosphatidylinositol 3-kinase (PI3K), a method of detecting a mutation in the gene encoding the regulatory p85.alpha. subunit of phosphatidylinositol 3-kinase, as well as a diagnostic composition and a test kit for use in the method.

20 Claims, 2 Drawing figures Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	KOMC
Drawi De	so Ir	nage							•	

4. Document ID: US 5246844 A

L9: Entry 4 of 4

File: USPT

Sep 21, 1993

US-PAT-NO: 5246844

DOCUMENT-IDENTIFIER: US 5246844 A

TITLE: Virulence associated proteins in Borrelia burgdorferi (BB)

DATE-ISSUED: September 21, 1993

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Norris; Steven J. Houston TX Barbour; Alan G. San Antonio TX

US-CL-CURRENT: $\frac{435}{480}$; $\frac{435}{252.3}$, $\frac{435}{252.3}$, $\frac{435}{252.33}$, $\frac{435}{320.1}$, $\frac{435}{476}$, $\frac{435}{488}$, $\frac{536}{23.7}$,

GB

WEST

Generate Collection

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Search Results - Record(s) 1 through 20 of 25 returned.

1. Document ID: US 20020137720 A1

L10: Entry 1 of 25

File: PGPB

Sep 26, 2002

PGPUB-DOCUMENT-NUMBER: 20020137720

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20020137720 A1

TITLE: Papilloma virus sequences

PUBLICATION-DATE: September 26, 2002

INVENTOR-INFORMATION:

Walcott, Sarah Marina

NAME CITY STATE COUNTRY RULE-47 Ertl, Peter F. Stevenage GB GB Gough, Gerald W. Stevenage GB Ring, Christopher Jeffrey Alan Stevenage GB Parmar, Vanita Stevenage

US-CL-CURRENT: 514/45; 435/235.1, 435/252.3, 435/325, 435/91.1, 514/44, 536/23.72

Full Title Citation Front Review Classification Date Reference Sequences Attachments Claims KMC Draw, Desc Image

Stevenage

2. Document ID: US 20020123083 A1

L10: Entry 2 of 25

File: PGPB

Sep 5, 2002

PGPUB-DOCUMENT-NUMBER: 20020123083

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20020123083 A1

TITLE: Nucleic acid endocing growth factor protein

PUBLICATION-DATE: September 5, 2002

INVENTOR-INFORMATION:

NAME CITY STATE COUNTRY RULE-47

Shigeta, Ron T. JR. Berkeley CA US Siani-Rose, Michael A. San Francisco CA US

US-CL-CURRENT: 435/7.23; 435/320.1, 435/325, 435/69.4, 530/399, 536/23.5, 702/19, 800/8

Full Title Citation Front Review Classification Date Reference Sequences Attachments Claims KWC Draw Desc Image

3. Document ID: US 6366860 B1

L10: Entry 3 of 25

File: USPT

Apr 2, 2002

US-PAT-NO: 6366860

DOCUMENT-IDENTIFIER: US 6366860 B1

TITLE: Synthetic genes for enhanced expression

DATE-ISSUED: April 2, 2002

INVENTOR - INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Rozzell, Jr.; J. David Burbank CA Bui; Peter El Monte CA

US-CL-CURRENT: 702/27; 540/456, 540/460, 702/19, 702/30, 702/32

ABSTRACT:

A method of making a synthetic nucleic acid sequence comprises providing a starting nucleic acid sequence, which optionally encodes an amino acid sequence, and determining the predicted .DELTA.G.sub.folding of the sequence. The starting nucleic acid sequence can be a naturally occurring sequence or a non-naturally occurring sequence. The starting nucleic acid sequence is modified by replacing at least one codon from the starting nucleic acid sequence with a different corresponding codon to provide a modified nucleic acid sequence. As used herein, a "different corresponding codon" refers to a codon which does not have the identical nucleotide sequence, but which encodes the identical amino acid. The predicted .DELTA.G.sub.folding of the modified nucleic acid sequence is determined and compared with the .DELTA.G.sub.folding of the starting nucleic acid sequence. In accordance with the invention, the predicted .DELTA.G.sub.folding of the starting nucleic acid sequence can be determined before or after the modified starting nucleic acid is provided.

23 Claims, 10 Drawing figures Exemplary Claim Number: 1 Number of Drawing Sheets: 7

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	KWIC
Draw, De	ese li	nage					- 4, 4,4,5			

4. Document ID: US 6365390 B1

L10: Entry 4 of 25 File: USPT Apr 2, 2002

US-PAT-NO: 6365390

DOCUMENT-IDENTIFIER: US 6365390 B1

TITLE: Phenolic acid esterases, coding sequences and methods

DATE-ISSUED: April 2, 2002

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY Blum; David L. San Diego CA Kataeva; Irina Athens GA Li; Xin-Liang Athens GA Ljungdahl; Lars G. Athens GA

US-CL-CURRENT: 435/197; 435/183, 435/252.3, 435/320.1, 530/350, 536/23.1, 536/23.2

ABSTRACT:

Described herein are four phenolic acid esterases, three of which correspond to domains of previously unknown function within bacterial xylanases, from XynY and XynZ of Clostridium thermocellum and from a xylanase of Ruminococcus. The fourth specifically exemplified xylanase is a protein encoded within the genome of Orpinomyces PC-2. The amino acids of these polypeptides and nucleotide sequences encoding them are provided. Recombinant host cells, expression vectors and methods for the recombinant production of phenolic acid esterases are also provided.

26 Claims, 13 Drawing figures Exemplary Claim Number: 1 Number of Drawing Sheets: 11

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments
Draw, D	esc li	mage							

5. Document ID: US 6337181 B1

L10: Entry 5 of 25

File: USPT

Jan 8, 2002

US-PAT-NO: 6337181

DOCUMENT-IDENTIFIER: US 6337181 B1

TITLE: Method of specifying vaccine components for viral quasispecies

DATE-ISSUED: January 8, 2002

INVENTOR-INFORMATION:

NAME CITY ZIP CODE COUNTRY STATE Stewart; Jeffrey Joseph N.T 07928 Chatham Elkins Pk. PΑ 19027 Litwin; Samuel Watts; Perry Elkins Pk. PA 19027

US-CL-CURRENT: $\underline{435/5}$; $\underline{424/184.1}$, $\underline{424/206.1}$, $\underline{424/93.21}$, $\underline{435/320.1}$, $\underline{435/325}$, $\underline{435/455}$, $\underline{514/44}$, $\underline{530/351}$

ABSTRACT:

An algorithm for determining the viral antigenic protein variants to be used to construct vaccines designed to immunize against variable viral populations (quasispecies) is described. The method entails analyzing multiple nucleotide sequences of viral proteins and identifying those variants that provide selective advantage to the virus. Examples are given for influenza A hemagglutinin 3 and HIV-1 gp120.

16 Claims, 2 Drawing figures Exemplary Claim Number: 1 Number of Drawing Sheets: 2

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	KWMC
Draw. D	esc	mage								

6. Document ID: US 6333172 B1

Record List Display

L10: Entry 6 of 25

File: USPT

Dec 25, 2001

US-PAT-NO: 6333172

DOCUMENT-IDENTIFIER: US 6333172 B1

TITLE: Genes and proteins controlling cholesterol synthesis

DATE-ISSUED: December 25, 2001

INVENTOR - INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Rine; Jasper D. Moraga CA Hampton; Randolph San Diego CA

US-CL-CURRENT: 435/69.1; 435/252.3, 435/320.1, 435/455

ABSTRACT:

The present invention provides isolated nucleic acid sequences which encode a family of HMG-CoA Reductase Degradation (HRD) polypeptides. More particularly, the present invention provides isolated HRD1, HRD2 and HRD3 nucleic acids and the Hrd polypeptides encoded by such nucleic acids, i.e., Hrd1, Hrd2 and Hrd3, respectively. Vectors comprising the nucleic acids are provided. In addition, the present invention provides screening assay related to cholesterol biosynthesis.

36 Claims, 11 Drawing figures Exemplary Claim Number: 1 Number of Drawing Sheets: 10

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments
Drawt D		mage							

KWIC

7. Document ID: US 6232458 B1

L10: Entry 7 of 25

File: USPT

May 15, 2001

US-PAT-NO: 6232458

DOCUMENT-IDENTIFIER: US 6232458 B1

TITLE: Synthetic polynucleotides encoding tropoelastin

DATE-ISSUED: May 15, 2001

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY Weiss; Anthony Steven Sydney AU Martin; Stephen Lewis Sedgley GB

US-CL-CURRENT: $\underline{536}/\underline{23.5}$; $\underline{435}/\underline{252.33}$, $\underline{435}/\underline{254.1}$, $\underline{435}/\underline{254.2}$, $\underline{435}/\underline{320.1}$, $\underline{435}/\underline{69.1}$, $\underline{435}/\underline{69.7}$, $\underline{530}/\underline{353}$, $\underline{536}/\underline{23.4}$, $\underline{536}/\underline{24.1}$, $\underline{536}/\underline{24.2}$

ABSTRACT:

Recombinant tropoelastins and variants of recombinant tropoelastins produced from synthetic polynucleotides, as well as the synthetic polynucleotides themselves are provided. Also provided are cross-linked elastins or elastin-like products prepared from the tropoelastins or variants.

24 Claims, 21 Drawing figures Exemplary Claim Number: 1

Number of Drawing Sheets: 21

Full Title Citation Front Review Classification Date Reference Sequences Attachments KMC |
Draw Desc | Image |

8. Document ID: US 6184018 B1

L10: Entry 8 of 25

File: USPT

Feb 6, 2001

US-PAT-NO: 6184018

DOCUMENT-IDENTIFIER: US 6184018 B1

TITLE: .beta.-glucosidase coding sequences and protein from orpinomyces PC-2

DATE-ISSUED: February 6, 2001

INVENTOR-INFORMATION:

ZIP CODE COUNTRY CITY STATE NAME Athens GA Li; Xin-Lianq GA Ljungdahl; Lars G. Athens Lawrenceville GA Chen; Huizhong Ximenes; Eduardo A. Athens GA

ABSTRACT:

Provided is a novel .beta.-glucosidase from Orpinomyces sp. PC2, nucleotide sequences encoding the mature protein and the precursor protein, and methods for recombinant production of this .beta.-glucosidase.

13 Claims, 7 Drawing figures Exemplary Claim Number: 1 Number of Drawing Sheets: 5



KWAC

9. Document ID: US 6114158 A

L10: Entry 9 of 25

File: USPT

Sep 5, 2000

US-PAT-NO: 6114158

DOCUMENT-IDENTIFIER: US 6114158 A

TITLE: Orpinomyces cellulase celf protein and coding sequences

DATE-ISSUED: September 5, 2000

INVENTOR-INFORMATION:

Record List Display

NAME CITY STATE ZIP CODE COUNTRY

Li; Xin-Liang Athens GA
Chen; Huizhong Athens GA
Ljungdahl; Lars G. Athens GA

US-CL-CURRENT: <u>435/209</u>; <u>435/252.3</u>, <u>536/23.2</u>

ABSTRACT:

A cDNA (1,520 bp), designated celF, consisting of an open reading frame (ORF) encoding a polypeptide (CelF) of 432 amino acids was isolated from a cDNA library of the anaerobic rumen fungus Orpinomyces PC-2 constructed in Escherichia coli. Analysis of the deduced amino acid sequence showed that starting from the N-terminus, CelF consists of a signal peptide, a cellulose binding domain (CBD) followed by an extremely Asn-rich linker region which separate the CBD and the catalytic domains. The latter is located at the C-terminus. The catalytic domain of CelF is highly homologous to CelA and CelC of Orpinomyces PC-2, to CelA of Neocallimastix patriciarum and also to cellobiohydrolase IIs (CBHIIs) from aerobic fungi. However, Like CelA of Neocallimastix patriciarum, CelF does not have the noncatalytic repeated peptide domain (NCRPD) found in CelA and CelC from the same organism. The recombinant protein CelF hydrolyzes cellooligosaccharides in the pattern of CBHII, yielding only cellobiose as product with cellotetraose as the substrate. The genomic celF is interrupted by a 111 bp intron, located within the region coding for the CBD. The intron of the celF has features in common with genes from aerobic filamentous fungi.

20 Claims, 3 Drawing figures Exemplary Claim Number: 1 Number of Drawing Sheets: 4

> িFull | Title | Citation |িFront | Review | Classification | Date | Reference | Sequences | Attachments | Draw Desc | Image |

KWMC

10. Document ID: US 6110720 A

L10: Entry 10 of 25

File: USPT

Aug 29, 2000

US-PAT-NO: 6110720

DOCUMENT-IDENTIFIER: US 6110720 A

TITLE: Orpinomyces cellulase CelE protein and coding sequences

DATE-ISSUED: August 29, 2000

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Li; Xin-Liang Athens GA
Ljungdahl; Lars G. Athens GA
Chen; Huizhong Athens GA

US-CL-CURRENT: 435/209; 435/252.3, 435/252.33, 536/23.2

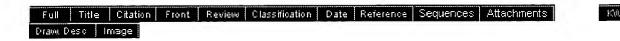
ABSTRACT:

A CDNA designated celE cloned from Orpinomyces PC-2 encodes a polypeptide (CelE) of 477 amino acids. CelE is highly homologous to CelB of Orpinomyces (72.3% identity) and Neocallimastix (67.9% identity), and like them, it has a non-catalytic repeated peptide domain (NCRPD) at the C-terminal end. The catalytic domain of CelE is homologous to glycosyl hydrolases of Family 5, found in several anaerobic bacteria. The gene of celE is devoid of introns. The recombinant proteins CelE and CelB of Orpinomyces PC-2 randomly hydrolyze carboxymethylcellulose and cello-oligosaccharides in the pattern of

ZIP CODE

endoglucanases.

9 Claims, 2 Drawing figures Exemplary Claim Number: 1 Number of Drawing Sheets: 2



11. Document ID: US 6107462 A

L10: Entry 11 of 25

File: USPT

Aug 22, 2000

COUNTRY

US-PAT-NO: 6107462

DOCUMENT-IDENTIFIER: US 6107462 A

TITLE: Genes and proteins controlling cholesterol synthesis

DATE-ISSUED: August 22, 2000

INVENTOR-INFORMATION:

NAME CITY STATE

Rine; Jasper D. Moraga CA Hampton; Randolph San Diego CA

US-CL-CURRENT: 530/350; 435/69.1, 536/23.5, 536/23.7

ABSTRACT:

The present invention provides isolated nucleic acid sequences which encode a family of HMG-CoA Reductase Degradation (HRD) polypeptides. More particularly, the present invention provides isolated HRD1, HRD2 and HRD3 nucleic acids and the Hrd polypeptides encoded by such nucleic acids, i.e., Hrd1, Hrd2 and Hrd3, respectively. Vectors comprising the nucleic acids are provided. In addition, the present invention provides screening assay related to cholesterol biosynthesis.

5 Claims, 10 Drawing figures Exemplary Claim Number: 1 Number of Drawing Sheets: 10

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments
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12. Document ID: US 5989903 A

L10: Entry 12 of 25

File: USPT

Nov 23, 1999

US-PAT-NO: 5989903

DOCUMENT-IDENTIFIER: US 5989903 A

TITLE: Strain for the production of 6-dimethyltetracycline, method for producing the

strain and vector for use in the method

DATE-ISSUED: November 23, 1999

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Ryan; Michael J.

West Milford

NJ

US-CL-CURRENT: $\frac{435}{320.1}$; $\frac{435}{183}$, $\frac{435}{252.3}$, $\frac{435}{252.35}$, $\frac{435}{471}$, $\frac{435}{64}$, $\frac{536}{23.2}$, $\frac{536}{23.7}$, $\frac{536}{24.1}$

ABSTRACT:

Recombinant S. aureofaciens cells are provided. These cells comprise:

- (a) at least one CTC 11 gene; and (b) optionally
- (i) a CTC 09 gene;
- (ii) a CTC 03 gene; or
- (iii) a combination thereof;

wherein:

the CTC 11 gene is chromosomal, extra-chromosomal, or chromosomal and extra-chromosomal;

the CTC 09 gene, CTC 03 gene, or a combination thereof is chromosomal, extra-chromosomal, or a combination thereof;

expression of the CTC 11 gene is enhanced over that of a wild-type S. aureofaciens cell; and

optionally, the CTC 09 gene, the CTC 03 gene, or both of the CTC 09 gene and the CTC 03 gene are inactivated.

The present invention also contemplates vector pLP21329 and vectors for allelic replacement in a S. aureofaciens host cell. The vectors comprise:

- (a) a functional E. coli origin of replication;
- (b) a functional Streptomyces origin of replication;
- (c) a functional gene that imparts a positively selectable phenotype on the host cell; and
- ((d) a ribosomal S12 gene which is expressed in Streptomyces such that it imparts sensitivity to streptomycin to the host cell.

In another embodiment, a method of mutating a target gene of a biosynthetic pathway of Streptomyces is disclosed. The method comprises

- (a) replacing the genomic copy of the target gene with a selectable marker gene through homologous recombination to form a first recombinant strain; and
- (b) replacing the selectable marker gene in the first recombinant strain with an altered copy of the target gene through homologous recombination to form a second recombinant strain.
- 12 Claims, 32 Drawing figures Exemplary Claim Number: 1 Number of Drawing Sheets: 47

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	KWAC
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13. Document ID: US 5986080 A

Record List Display

L10: Entry 13 of 25

File: USPT

Nov 16, 1999

US-PAT-NO: 5986080

DOCUMENT-IDENTIFIER: US 5986080 A

TITLE: Cloned nucleotide pyrophosphohydrolase and uses thereof

DATE-ISSUED: November 16, 1999

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Masuda; Ikuko	Wauwatosa	WI		
Barbieri; Joseph T.	New Berlin	WI		
Haas; Arthur L.	Brookfield	WI		
Halligan; Brian D.	Wauwatosa	WI		
McCarty; Daniel J.	Hartland	WI		
Ryan; Lawrence M.	Wauwatosa	WI		

US-CL-CURRENT: 536/23.2; 435/195, 435/252.3, 435/320.1, 530/350

ABSTRACT:

We have cloned and sequenced the cDNA encoding the 61 kD active fragment of a unique porcine chondrocyte nucleotide pyrophosphohydrolase (NTPPHase) from a porcine chondrocyte library. Degenerate oligonucleotides, corresponding to the N-terminal amino acid sequence of this peptide were hybridized to porcine chondrocyte cDNA and used to amplify DNA encoding the N-terminal sequence of 61 kD with the polymerase chain reaction (PCR). The PCR products were then used as probes to clone the entire open reading-frame for the 61 kD fragment from a porcine chondrocyte cDNA library. The length of the cloned cDNA was 2509 bp. Translation of the open-reading-frame predicts the 61 kD fragment to be a 459 amino acid protein. BLAST and FASTA analysis confirmed that this amino acid sequence was unique and did not possess high homology to any known proteins in the non-redundant protein data base. Limited homology (17%) between the 61 kD fragment and several prokaryotic and eukaryotic ATP pyrophosphate-lyase (adenylate cyclase) was detected. Northern blot analysis of porcine chondrocyte RNA showed that the DNA encoding the 61 kD fragment hybridized to a 4.3 kbp RNA transcript. Human chondrocyte RNA also hybridized to this porcine DNA probe. Coupled in vitro transcription translation of an expression vector containing the DNA insert in frame showed the expression of a 61 kD protein.

3 Claims, 9 Drawing figures Exemplary Claim Number: 1 Number of Drawing Sheets: 9

Full | Title | Citation | Front | Review | Classification | Date | Reference | Sequences | Attachments | Draw, Desc | Image |

KWIC

14. Document ID: US 5965429 A

L10: Entry 14 of 25

File: USPT

Oct 12, 1999

US-PAT-NO: 5965429

DOCUMENT-IDENTIFIER: US 5965429 A

TITLE: Strain for the production of 6-demethyltetracycline, method for producing the strain and vector for use in the method

betain and vector for abe in the met

DATE-ISSUED: October 12, 1999

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Ryan; Michael J. West Milford NJ

US-CL-CURRENT: $\frac{435}{252.35}$; $\frac{435}{253.5}$, $\frac{435}{320.1}$, $\frac{435}{64}$, $\frac{435}{889}$, $\frac{536}{23.1}$

ABSTRACT:

Recombinant S. aureofaciens cells are provided. These cells comprise: (a) at least one CTC 11 gene; and (b) optionally

- (i) a CTC 09 gene;
- (ii) a CTC 03 gene; or
- (iii) a combination thereof;

wherein:

the CTC 11 gene is chromosomal, extra-chromosomal, or chromosomal and ${\sf extra-chromosomal}$;

the CTC 09 gene, CTC 03 gene, or a combination thereof is chromosomal, extra-chromosomal, or a combination thereof; expression of the CTC 11 gene is enhanced over that of a wild-type S. aureofaciens cell; and

optionally, the CTC 09 gene, the CTC 03 gene, or both of the CTC 09 gene and the CTC 03 gene are inactivated.

The present invention also contemplates vector pLP21329 and vectors for allelic replacement in a S. aureofaciens host cell. The vectors comprise:

- (a) a functional E. coli origin of replication;
- (b) a functional Streptomyces origin of replication;
- (c) a functional gene that imparts a positively selectable phenotype on the host cell; and
- ((d) a ribosomal S12 gene which is expressed in Streptomyces such that it imparts sensitivity to streptomycin to the host cell. In another embodiment, a method of mutating a target gene of a biosynthetic pathway of Streptomyces is disclosed. The method comprises
- (a) replacing the genomic copy of the target gene with a selectable marker gene through homologous recombination to form a first recombinant strain; and
- (b) replacing the selectable marker gene in the first recombinant strain with an altered copy of the target gene through homologous recombination to form a second recombinant strain.

27 Claims, 48 Drawing figures Exemplary Claim Number: 1 Number of Drawing Sheets: 47

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	KWAC
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15. Document ID: US 5955277 A

L10: Entry 15 of 25 File: USPT Sep 21, 1999

US-PAT-NO: 5955277

DOCUMENT-IDENTIFIER: US 5955277 A

Record List Display

TITLE: Mutant cDNA encoding the p85.alpha. subunit of phosphatidylinositol 3-kinase

DATE-ISSUED: September 21, 1999

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Hansen; Torben Hellerup DK

Andersen; Carsten Bo Los Altos CA

Pedersen; Oluf Borbye Holte DK

US-CL-CURRENT: 435/6; 435/91.2, 536/23.1

ABSTRACT:

The present invention relates to a mutant cDNA sequence encoding the regulatory p85.alpha. subunit of phosphatidylinositol 3-kinase (PI3K), a method of detecting a mutation in the gene encoding the regulatory p85.alpha. subunit of phosphatidylinositol 3-kinase, as well as a diagnostic composition and a test kit for use in the method.

20 Claims, 2 Drawing figures Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments
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KMC

16. Document ID: US 5945292 A

L10: Entry 16 of 25 File: USPT Aug 31, 1999

US-PAT-NO: 5945292

DOCUMENT-IDENTIFIER: US 5945292 A

TITLE: Method of identifying cells with polypeptide surface marker

DATE-ISSUED: August 31, 1999

INVENTOR - INFORMATION:

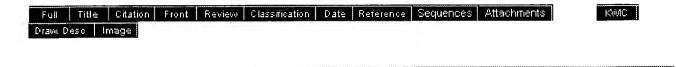
CITY NAME STATE ZIP CODE COUNTRY Brizzard; Billy L. New Haven CTBianca; Darlene W. Westbrook CTChubet; Richard G. Middletown Vizard; Douglas L. Cheshire CTHopp; Thomas Patrick San Diego CA

US-CL-CURRENT: 435/7.21; 435/29, 435/34

ABSTRACT:

This invention discloses a gene for the identification of cells comprising a secretion leader segment, a cell marker segment and a transmembrane segment. The gene can be used to identify cells transfected with the gene by the steps of: inserting the gene having a secretion leader segment, a cell marker segment and a transmembrane segment into the DNA or RNA of a cell, allowing the cell to express the gene, and detecting the expressed cell marker segment of the gene.

10 Claims, 4 Drawing figures Exemplary Claim Number: 1 Number of Drawing Sheets: 3



17. Document ID: US 5731425 A

L10: Entry 17 of 25

File: USPT

Mar 24, 1998

US-PAT-NO: 5731425

DOCUMENT-IDENTIFIER: US 5731425 A

TITLE: Polypeptide surface marker for cells

DATE-ISSUED: March 24, 1998

INVENTOR-INFORMATION:

CITY STATE ZIP CODE COUNTRY NAME CTBrizzard; Billy L. New Haven Bianca; Darlene W. Westbrook CTCTChubet; Richard G. Middletown Vizard; Douglas L. Cheshire CTCA Hopp; Thomas Patrick San Diego

US-CL-CURRENT: 536/23.1; 435/320.1, 435/69.1, 536/24.1

ABSTRACT:

This invention discloses a gene for the identification of cells comprising a selection leader segment, a cell marker segment and a transmembrane segment. The gene can be used to identify cells transfected with the gene by the steps of: inserting the gene having a selection leader segment, a cell marker segment and a transmembrane segment into the DNA or RNA of a cell, allowing the cell to express the gene, and detecting the expressed cell marker segment of the gene.

12 Claims, 4 Drawing figures Exemplary Claim Number: 1 Number of Drawing Sheets: 3

Full Tit	le	Citation	Front	Review	Classification		Reference	Sequences	
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18. Document ID: US 5677153 A

L10: Entry 18 of 25

File: USPT

Oct 14, 1997

US-PAT-NO: 5677153

DOCUMENT-IDENTIFIER: US 5677153 A

TITLE: Methods for modifying DNA and for detecting effects of such modification on interaction of encoded modified polypeptides with target substrates

DATE-ISSUED: October 14, 1997

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Botstein; David Belmont CA Palzkill; Timothy Union City CA

US-CL-CURRENT: 435/91.4; 435/91.2, 435/91.41, 435/91.42

ABSTRACT:

The invention relates to methods and mutation linkers to modify DNA, to methods for producing libraries containing a multiplicity of modified DNA, and to methods for using such libraries for screening modified proteins encoded by such DNA. The DNA targeted for modification typically encodes a polypeptide such as an enzyme. The libraries are used to determine the effect of such modification or the interaction of the modified polypeptides with a target. In preferred embodiments, the invention relates to methods for making and using libraries containing DNA encoding modified antibiotic hydrolases to screen antibiotics against one or more of the modified antibiotic hydrolases produced by such libraries. Susceptibility or lack of susceptibility of an antibiotic to neutralization provides an indication of whether wild-type antibiotic hydrolases are likely to mutate to confer resistance to the antibiotic.

40 Claims, 55 Drawing figures Exemplary Claim Number: 2 Number of Drawing Sheets: 23

Full Title Citation Front Review Classification Date Reference Sequences Attachments
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19. Document ID: US 5670356 A

L10: Entry 19 of 25 File: USPT Sep 23, 1997

US-PAT-NO: 5670356

DOCUMENT-IDENTIFIER: US 5670356 A

TITLE: Modified luciferase

DATE-ISSUED: September 23, 1997

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

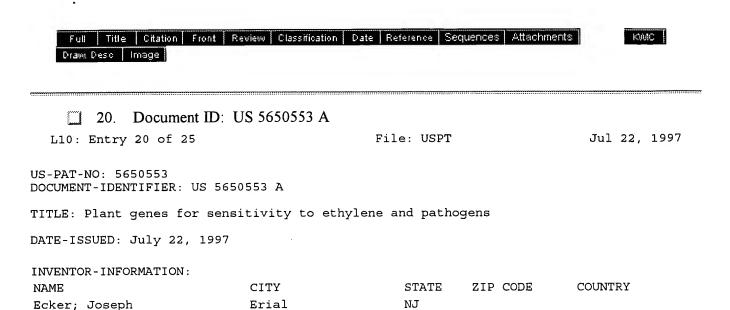
Sherf; Bruce A. Waunakee WI Wood; Keith V. Madison WI

US-CL-CURRENT: 435/189; 435/358, 435/364, 435/367, 435/394, 435/455, 536/23.2

ABSTRACT:

A modified form of beetle luciferase, which has been engineered for improved genetic reporting, is disclosed. The modified form contains one or more new features. Chief among these is removal of the peroxisomal translocation sequence to yield a cytoplasmic form of the enzyme. Other changes include removal of potentially interfering restriction sites and genetic regulatory sites from the gene, improvement of the codon usage for mammalian cells. The modified luciferase reporter enzyme is also devoid of potential N-glycosylation targets to minimize post-translational modification and remains in the cytoplasm of host cells to optimize substrate availability.

17 Claims, 6 Drawing figures Exemplary Claim Number: 1 Number of Drawing Sheets: 6



Lehman; Anne Philadelphia PA
Roman; Gregg North Wales PA

Haverford

US-CL-CURRENT: 800/298; 435/419, 514/12, 530/370, 536/23.6, 800/301

ABSTRACT:

Rothenberg; Madge

The present invention is directed to nucleic acid sequences for ethylene insensitive, EIN loci and corresponding amino acid sequences. The present invention is also directed to nucleic acid sequences for hookless1, HLS1, alleles and amino acid sequences.

PA

17 Claims, 42 Drawing figures Exemplary Claim Number: 16 Number of Drawing Sheets: 35

Full	Title	Citation	Front	Review	Classification		Reference	Sequences	Attachments	KWC
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Search Results - Record(s) 21 through 25 of 25 returned.

21. Document ID: US 5246844 A

L10: Entry 21 of 25

File: USPT

Sep 21, 1993

US-PAT-NO: 5246844

DOCUMENT-IDENTIFIER: US 5246844 A

TITLE: Virulence associated proteins in Borrelia burgdorferi (BB)

DATE-ISSUED: September 21, 1993

INVENTOR - INFORMATION:

NAME

CITY

STATE

ZIP CODE

COUNTRY

Norris; Steven J. Barbour; Alan G. Houston San Antonio TX TX

US-CL-CURRENT: $\frac{435}{480}$; $\frac{435}{252.3}$, $\frac{435}{252.33}$, $\frac{435}{320.1}$, $\frac{435}{476}$, $\frac{435}{488}$, $\frac{536}{23.7}$, $\frac{536}{24.32}$, $\frac{536}{24.33}$

ABSTRACT:

The invention relates to a DNA segment encoding a Borrelia burgdorferi antigenic polypeptide. The invention also relates to a purified 30 kDa polypeptide isolated from a virulent strain of B. burgdorferi and to epitopic segments of the polypeptide with immunogenic potential. The 30 kDa protein provides a route for the development of immunodiagnostics for Lyme disease and related disorders. The 30 kDa protein and related amino acid and DNA sequences may also be used for the immunization, for the detection of B. burgdorferi in human or animal tissues or body fluids, and also for the generation of specific antibodies for use in diagnosis, epidemiology, and prevention of Lyme disease.

22 Claims, 10 Drawing figures Exemplary Claim Number: 1 Number of Drawing Sheets: 14

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC
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22. Document ID: US 5082767 A

L10: Entry 22 of 25

File: USPT

Jan 21, 1992

US-PAT-NO: 5082767

DOCUMENT-IDENTIFIER: US 5082767 A

TITLE: Codon pair utilization

DATE-ISSUED: January 21, 1992

INVENTOR-INFORMATION:

NAME

CITY

STATE ZIP CODE COUNTRY

Hatfield; G. Wesley

Corona Del Mar

CA

92625

Gutman; George A.

Costa Mesa

92626

US-CL-CURRENT: 435/6; 435/69.1, 435/91.5, 436/501, 536/23.1, 536/24.1

ABSTRACT:

A method for determining the pattern of nonrandom codon pair usage of an organism, comprising the steps of obtaining nucleotide sequence data for the organism, determining from the data the number of codons represented in at least a portion of the sequence and the frequency of usage of at least some codons in the portion, determining from the frequency the expected number of occurrences of at least some codon pairs, if they are paired in a random manner, and comparing the expected number with the actual number of occurrences to determine relative codon pairing preferences. The codon pairings of organisms are highly nonrandom, and differ from organism to organism. This information is used to construct and express altered or synthetic genes having desired levels of translational efficiency, to determine which regions in a genome are protein coding regions, to introduce translational pause sites into heterologous genes, and to ascertain relationship or ancestral origin of nucleotide sequences.

44 Claims, 7 Drawing figures Exemplary Claim Number: 1 Number of Drawing Sheets: 4

> Full Title Citation Front Review Classification Date Reference Sequences Attachments Drawi Desc Image

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23. Document ID: WO 2064799 A2

L10: Entry 23 of 25

File: EPAB

Aug 22, 2002

PUB-NO: WO002064799A2

DOCUMENT-IDENTIFIER: WO 2064799 A2 TITLE: OPTIMIZED MESSENGER RNA

PUBN-DATE: August 22, 2002

INVENTOR - INFORMATION:

NAME SELDON, RICHARD F

MILLER, ALLAN M

TRECO, DOUGLAS S

COUNTRY

US

US

US

INT-CL (IPC): C12 N 15/67; C07 H 21/00; C07 K 14/745; C12 N 15/63

EUR-CL (EPC): C12N009/40; C07K014/745, C12N015/67

ABSTRACT:

The present invention is directed to a synthetic nucleic acid sequence which encodes a protein wherein at least one non-common codon or less-common codon is replaced by a common codon. The synthetic nucleic acid sequence can include a continuous stretch of at least 90 codons all of which are common codons.

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments
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KWIC

24. Document ID: AU 200177554 A DE 10037111 A1 WO 200210411 A2

L10: Entry 24 of 25 File: DWPI Feb 13, 2002

DERWENT-ACC-NO: 2002-189232

DERWENT-WEEK: 200238

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TITLE: Preparing heterologous protein in prokaryotes, useful particularly for human growth hormone, with the coding sequence optimized for codon usage of the host

INVENTOR: BERGEMANN, K; GOETZ, F; PESCHEL, A; WERNER, R

PRIORITY-DATA: 2000DE-1037111 (July 27, 2000)

PATENT-FAMILY:

PUB-NO	PUB-DATE	LANGUAGE	PAGES	MAIN-IPC
AU 200177554 A	February 13, 2002		000	C12N015/67
DE 10037111 A1	February 7, 2002		016	C07H021/00
WO 200210411 A2	February 7, 2002	E	000	C12N015/67

INT-CL (IPC): $\underline{\text{CO7}}$ $\underline{\text{H}}$ $\underline{\text{21}}/\underline{\text{00}}$; $\underline{\text{C12}}$ $\underline{\text{N}}$ $\underline{\text{1}}/\underline{\text{21}}$; $\underline{\text{C12}}$ $\underline{\text{N}}$ $\underline{\text{15}}/\underline{\text{63}}$; $\underline{\text{C12}}$ $\underline{\text{N}}$ $\underline{\text{15}}/\underline{\text{67}}$; $\underline{\text{C12}}$ $\underline{\text{P}}$ $\underline{\text{21}}/\underline{\text{00}}$; $\underline{\text{C12}}$ $\underline{\text{P}}$ $\underline{\text{C12}}/\underline{\text{P}}$ $\underline{\text{C12}}/\underline{\text{P}}/\underline{\text{P}}$ $\underline{\text{C12}}/\underline{\text{P}}/\underline{\text{P}}$ $\underline{\text{C12}}/\underline{\text{P}}/\underline{\text{P}}/\underline{\text{P}}$ $\underline{\text{C12}}/\underline{\text{P}}/\underline{\text$

ABSTRACTED-PUB-NO: DE 10037111A

BASIC-ABSTRACT:

NOVELTY - Preparation of a heterologous recombinant protein (I) in a prokaryotic host cell (A) in which:

- (i) codon utilization of the host, for its own genes, is determined;
- (ii) the nucleic acid (II) that encodes (I) is modified to replace rare codons (for the host) by common codons; and
- (iii) cells are transformed with the modified sequence (IIa), for expression, is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- (a) similar method in which the host cell is transformed with nucleic acid encoding tRNA for rare codons;
- (b) (IIa) in which at least one rare codon has been replaced by a common codon;
- (c) a nucleic acid sequence (IIb) of about 1380 base pairs as given in the specification that encodes human growth hormone (hGH), also its fragments, sequences that hybridize to it under stringent conditions, its allelic or functional variants, and its variants within the degeneracy of the genetic code;
- (d) vectors containing (IIa) or (IIb);
- (e) host cells containing (IIa), (IIb) or the vectors of (d); and
- (f) host cells containing one or more tRNA corresponding to codons rarely used by the cells.

USE - The method is used for production of antibodies, insulin, tissue plasminogen activator and particularly human growth hormone.

ADVANTAGE - Codon optimization and/or incorporation of tRNA for rare codons significantly increases expression rate of (I), and thus the yield, as less proteolysis occurs.

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments
Draw, D	esc li	mage							

KWAC

25. Document ID: AU 200178641 A WO 200204494 A2

L10: Entry 25 of 25

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Jan 21, 2002

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TITLE: Selecting HIV-1 subtype C isolates, which are useful in developing vaccines against HIV infection, comprises isolating viruses with high sequence identity to a consensus sequence whose phenotype is associated with the HIV subtype

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PATENT-FAMILY:

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ABSTRACTED-PUB-NO: WO 200204494A

BASIC-ABSTRACT:

NOVELTY - Selecting human immunodeficiency virus (HIV) subtype isolates for use in the development of a prophylactic and/or therapeutic pharmaceutical composition, comprising selecting isolated virus or viruses with a high sequence identity to a consensus sequence, a phenotype which is associated with transmission for the particular HIV subtype, is new.

DETAILED DESCRIPTION - Selecting human immunodeficiency virus (HIV) subtype isolates for use in the development of a prophylactic and/or therapeutic pharmaceutical composition, comprising:

- (a) isolating viruses from recently infected subjects;
- (b) generating a consensus sequence for at least part of at least one HIV gene by identifying the most <u>common codon</u> or amino acid among the isolated viruses at each position along at least part of the gene; and
- (c) selecting the isolated virus or viruses with a high sequence identity to the consensus sequence, a phenotype that is associated with transmission for the particular HIV subtype.

INDEPENDENT CLAIMS are also included for the following:

- (1) HIV-1 subtype C isolates, which are designated Du422 (Provisional Accession Number 01032114, European Collection of Cell Cultures), Du151 (European Collection of Cell Cultures Accession Number 00072724) and Du179 (European Collection of Cell Cultures Accession Number 00072725);
- (2) molecules comprising the nucleic acid sequences of the sequenced gag gene of the isolate Du422, pol gene of the isolate Du151 or env gene of the isolate Du151, where the sequences are not fully defined in the specification; or a 2579 base pair sequence, fully defined in the specification;
- (3) polypeptides comprising the amino acid sequence of the sequenced gag gene of isolate Du422, pol gene of the isolate Du151 or env gene of the isolate Du151, where the sequences are not defined in the specification, an 858 residue amino acid sequence, fully defined in the specification, or a sequence that is a modification or derivative of them; and

- (4) consensus amino acid sequence for the partial:
- (a) gag gene of HIV-1 subtype C having a 313 residue amino acid sequence, fully defined in the specification;
- (b) pol gene of HIV-1 subtype C having a 278 residue amino acid sequence, fully defined in the specification; or
- (c) env gene of HlV-1 subtype C having a 229 residue amino acid sequence, fully defined in the specification.

ACTIVITY - Antiviral.

No biological data is given.

MECHANISM OF ACTION - Vaccine.

USE - For selecting HIV-1 subtype C isolates, which are useful in the development of a prophylactic and/or therapeutic pharmaceutical compositions, e.g. vaccines against HIV infection or disease.

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